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 NEWS  2  Apr 08  "Ask CAS" for self-help around the clock
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 NEWS  4  Apr 09  ZDB will be removed from STN
 NEWS  5  Apr 19  US Patent Applications available in IFICDB, IFIPAT, and IFIUDB
 NEWS  6  Apr 22  Records from IP.com available in CAPLUS, HCAPLUS, and ZCAPLUS
 NEWS  7  Apr 22  BIOSIS Gene Names now available in TOXCENTER
 NEWS  8  Apr 22  Federal Research in Progress (FEDRIP) now available
 NEWS  9  Jun 03  New e-mail delivery for search results now available
 NEWS 10  Jun 10  MEDLINE Reload
 NEWS 11  Jun 10  PCTFULL has been reloaded
 NEWS 12  Jul 02  FOREGE no longer contains STANDARDS file segment
 NEWS 13  Jul 22  USAN to be reloaded July 28, 2002;
                  saved answer sets no longer valid
 NEWS 14  Jul 29  Enhanced polymer searching in REGISTRY
 NEWS 15  Jul 30  NETFIRST to be removed from STN
 NEWS 16  Aug 08  CANCERLIT reload
 NEWS 17  Aug 08  PHARMAMarketLetter(PHARMAML) - new on STN
 NEWS 18  Aug 08  NTIS has been reloaded and enhanced
 NEWS 19  Aug 19  Aquatic Toxicity Information Retrieval (AQUIRE)
                  now available on STN
 NEWS 20  Aug 19  IFIPAT, IFICDB, and IFIUDB have been reloaded
 NEWS 21  Aug 19  The MEDLINE file segment of TOXCENTER has been reloaded
 NEWS 22  Aug 26  Sequence searching in REGISTRY enhanced
 NEWS 23  Sep 03  JAPIO has been reloaded and enhanced
 NEWS 24  Sep 16  Experimental properties added to the REGISTRY file
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 NEWS 26  Sep 16  CA Section Thesaurus available in CAPLUS and CA
 NEWS 27  Oct 01  CASREACT Enriched with Reactions from 1907 to 1985
 NEWS 28  Oct 21  EVENTLINE has been reloaded
 NEWS 29  Oct 24  BEILSTEIN adds new search fields
 NEWS 30  Oct 24  Nutraceuticals International (NUTRACEUT) now available on STN
 NEWS 31  Oct 25  MEDLINE SDI run of October 8, 2002                    on STN

 NEWS EXPRESS  October 14 CURRENT WINDOWS VERSION IS V6.01,
               CURRENT MACINTOSH VERSION IS V6.0a(ENG) AND V6.0Ja(JP),
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=> s testosterone (3W) hydroxylase
  37 FILES SEARCHED...
  81 FILES SEARCHED...
L1        4477 TESTOSTERONE (3W) HYDROXYLASE

=> s l1 (3A) p450
  42 FILES SEARCHED...
L2         237 L1 (3A) P450

=> s p450 (4A) cyp3a4
  49 FILES SEARCHED...
L3        2150 P450 (4A) CYP3A4

=> s l2 or l3
  20 FILES SEARCHED...
  61 FILES SEARCHED...
L4        2386 L2 OR L3

=> s (hepg2 (3a) transfection) and l4
  36 FILES SEARCHED...
  83 FILES SEARCHED...
L5           0 (HEPG2 (3A) TRANSFECTION) AND L4

=> s human (A) (liver or hepatocyte)
  13 FILES SEARCHED...
  23 FILES SEARCHED...
  38 FILES SEARCHED...
  47 FILES SEARCHED...
  68 FILES SEARCHED...
L6      122169 HUMAN (A) (LIVER OR HEPATOCYTE)

=> s l6 (3A) (cell line)
  11 FILES SEARCHED...
  13 FILES SEARCHED...
  26 FILES SEARCHED...
  30 FILES SEARCHED...
  42 FILES SEARCHED...
  52 FILES SEARCHED...
  59 FILES SEARCHED...
  72 FILES SEARCHED...
L7        3641 L6 (3A) (CELL LINE)

=> s l7 and l4
  47 FILES SEARCHED...
 75% OF LIMIT FOR L#S REACHED
L8           1 L7 AND L4 - test hydrox / P450 / CYP3A4
                           human liver / hepatocyte / cell line

=> d l8 bib ab

L8   ANSWER 1 OF 1  CAPLUS  COPYRIGHT 2002 ACS
AN   1995:971826  CAPLUS
DN   124:75473
TI   An investigation of the interaction between halofantrine, CYP2D6 and
     CYP3A4: Studies with human liver microsomes and heterologous enzyme
     expression systems
AU   Halliday, Rachel C.; Jones, Barry C.; Smith, Dennis A.; Kitteringham, Neil
     R.; Park, B. Kevin
```

CS Department Pharmacology and Therapeutics, University Liverpool, Liverpool, L69 3BX, UK
SO British Journal of Clinical Pharmacology (1995), 40(4), 369-78
CODEN: BCPHBM; ISSN: 0306-5251
PB Blackwell
DT Journal
LA English
AB The authors have assessed the interaction of the antimalarial halofantrine with cytochrome P 450 (CYP) enzymes in vitro, with the use of microsomes from **human liver** and recombinant **cell lines**. Rac-halofantrine was a potent inhibitor (IC50 = 1.06 .mu.M, Ki = 4.3 .mu.M) of the 1-hydroxylation of bufuralol, a marker for CYP2D6 activity. Of a group of structurally related antimalarials tested, only quinidine (IC50 = 0.04 .mu.M) was more potent. Microsomes prepd. from recombinant CYP2D6 and CYP3A4 cell lines were shown to catalyze halofantrine N-debutylation. The metab. of halofantrine to its N-desbutyl metabolite by human liver microsomes showed no correlation with CYP2D6 genotypic or phenotypic status and there was no consistent inhibition by quinidine. The rate of halofantrine metab. showed a significant correlation with both CYP3A4 protein levels (r = 0.88) and the rate of felodipine metab. (r = 0.86), a marker substrate for CYP3A4 activity. Inhibition studies showed that ketoconazole is a potent inhibitor of halofantrine metab. (IC50 = 1.57 .mu.M). In conclusion, the authors have demonstrated that halofantrine is a potent inhibitor of CYP2D6 in vitro and can also be metabolized by the enzyme. However, in human liver microsomes it appears to be metabolized largely by CYP3A4.

=>
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<------User Break------>

=> s transfection (3a) (l7 or hepg2)
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COMMAND INTERRUPTED
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  85 FILES SEARCHED...
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=> s l7 or hepg2
  48 FILES SEARCHED...
  92 FILES SEARCHED...
L9   52051 L7 OR HEPG2

=> s transfection (3a) l9
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  43 FILES SEARCHED...
  90 FILES SEARCHED...
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=> s l9 and p450
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 95% OF LIMIT FOR L#S REACHED
L10        2262 L9 AND P450

=> s l10 and transfection
MAXIMUM L# WOULD BE EXCEEDED.

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